

**In the specification**

Please replace the paragraph on page 35, lines 8-31 with the following paragraph.

The cloned p35 cDNA, after sequence confirmation, was adapted for expression as a fusion protein as follows. At the fusion junction, the C-terminal amino acid residue of the CH3 is lysine and the N-terminal residue of the mature p35 is arginine. To minimise proteolysis at the fusion junction with these two basic residues, both of them were mutagenised to alanine, which, in the case of IL2 immunocytokine, has been shown to extend serum half-life [Gillies *et al.* (2002) *Clin. Cancer Res.* **8**:210]. For reconstruction of the fusion junction, there is a convenient *B*alI site just 11 base-pairs (bp) downstream of the mature N-terminus of p35. Hence a *Xma*I–*B*alI oligonucleotide linker consisting of sense strand 5'-CCG GGC **GCC GCA AAC CTC CCC GTG G-3'** [SEQ ID NO:22] and anti-sense strand 5'- C CAC GGG GAG GTT **TGC GGC GC-3'** [SEQ ID NO:23], where the **GCC GCA** [SEQ ID NO: 24] denote the two alanine substitutions, was synthesised and ligated to a *B*alI-XhoI restriction fragment encoding the rest of the p35 subunit. The resultant *Xma*I-XhoI fragment in turn was ligated to the unique *Xma*I site in the pdHL11 expression vector, forming the CH3-p35 fusion junction. The peptide sequence at the junction, LSLSPGAANLPV [SEQ ID NO: 25] [SEQ ID NO:24], where AA are the two alanine substitutions, is novel and potentially immunogenic. Indeed it contained a potential T helper cell epitope, which could be removed by mutating the LSLS [SEQ ID NO: 26] residues to ATAT [SEQ ID NO: 27], based on Biovation's technology called deImmunization. The resultant deImmunised fusion junction sequence is called M1. Therefore, the huBC1-H chain-M1-hu p35 DNA consists of the XhoI-HindIII fragment encoding the signal peptide-VH,

SUPPLEMENTAL PRELIMINARY AMENDMENT AND TRANSMITTAL OF SEQUENCE LISTING

the HindIII-XmaI fragment encoding the genomic human IgG1 H chain constant regions with the deImmunised junction, and the XmaI-XhoI fragment encoding the p35 subunit.

Please replace the paragraph on page 46, 1-2 with the following paragraph.

PEHFSGRPREDRVPHSRNSITLTNLTPGTEYVVSIVAL

NGREESPLLIGRSRSHHHHHH"

[SEQ ID NO: 28][SEQ ID

**NO: 25}**

Please replace the paragraph on page 46, lines 5-9 with the following paragraph.

Note1: Residue 1 to 207 is pQE sequence from and including Qiagen promoter primer (CCCGAAAAGTGCCACCTG) [SEQ ID NO:29]. Residue 1069 to 1126 is pQE12 sequence from the end of the hexa-histidine tag to the Qiagen reverse primer sequence (GTTCTGAGGTCAATTACTGG) [SEQ ID NO:30]. Fibronectin-derived sequence (*i.e.* without MRGS and hexa-histidine tag is in lower case).

Please replace the paragraph on page 46, line 18 until page 47, line 4 with the following paragraph.

1 CCCCGAAAAG TGCCACCTGA CGTCTAAGAA ACCATTATTA TCATGACATT  
AACCTATAAA

61 AATAGGCGTA TCACGAGGCC CTTCGTCTT CACCTCGAGA AATCATAAAA  
AATTTATTTG

121 CTTTGTGAGC GGATAACAAT TATAATAGAT TCAATTGTGA GCGGATAACA  
ATTCACACA

SUPPLEMENTAL PRELIMINARY AMENDMENT AND TRANSMITTAL OF SEQUENCE LISTING

181 GAATTCACTTA AAGAGGAGAA ATTAACATATG AGAGGATCtg tggtagacacc  
attgtctcca  
241 ccaacaaact tgcacatcgga ggcaaaccct gacactggag tgctcacagt ctccctgggag  
301 aggaggacca cccccagacat tactggttat agaattacca caacccctac aaacggccag  
361 cagggaaatt ctttggaga agtggtccat gctgatcaga gctccgcac ttttgataac  
421 ctgagtcgg gcctggagta caatgtcagt gtttacactg tcaaggatga caaggaaagt  
481 gtccttatct ctgataccat catcccagct gttccctcctc ccactgaccc tcgcattcacc  
541 aacattggtc cagacaccat gcgtgtcacc tgggctccac ccccatccat tgatttaacc  
601 aactccctgg tgcgttactc acctgtgaaa aatgaggaag atgtgcaga gttgtcaatt  
661 ttccttcag acaatgcagt ggtcttaaca aatctcctgc ctggtagaca atatgttagtg  
721 agtgtctcca gtgtctacga acaacatgag agcacaccc tttagaggaag acagaaaaaca  
781 ggtcttgatt ccccaactgg cattgacttt tctgatatta ctgccaactc ttttactgtg  
841 cactggattt ctccctcgagc caccatcaact ggctacagga tccgcacatca tcccgagcac  
901 ttcagtggga gacctcgaga agatcggttg ccccaactctc ggaattccat caccctcacc  
961 aacccactc caggcacaga gtatgtggtc agcatcggtt ctcttaatgg cagagaggaa  
1021 agtcccttat tgattggcaG ATCCAGATCT CATCACCATC ACCATCACTA  
AGCTTAATTA

1081 GCTGAGCTTG GACTCCTGTT GATAGATCCA GTAATGACCT CAGAAC

[SEQ ID NO: 31] {SEQ ID NO: 26}

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SUPPLEMENTAL PRELIMINARY AMENDMENT AND TRANSMITTAL OF SEQUENCE LISTING

Please replace the paragraph on page 47, lines 6-36, with the following paragraph.

(b) *Fibronectin 7B89 fragment*

LOCUS FN7B89.DNA 1399 bp mRNA PRI 01-OCT-1999

DEFINITION Human mRNA for fibronectin domains 7B89 in pQE12 (pAS33)

NID Derived from g31396 and pQE12 (Qiagen).

VERSION X02761.1 GI:31396

KEYWORDS alternate splicing; fibronectin.

SOURCE human.

ORGANISM Homo sapiens

Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Mammalia;

Eutheria; Primates; Catarrhini; Hominidae; Homo.

CDS <208..1341

/product="Fn MRGS-7B89-HHHHHH"

/translation="

MRGSVVTPSLSPPTNLHLEANPDTGVLTVSWERSTTPDI

TGYRITTTPTNGQQGNSLEEVVHADQSSCTFDNLSPGL

EYNVSVYTVKDDKESVPISDTIPEVPQLTDLSFVDIT

DSSIGLRWTPLNSSTIIGYRITVVAAGEGIPIFEDFVD

SSVGYYTGTGLEPGIDYDISVITLINGGESAPTLTQQ

TAVPPPDTDLRFTNIGPDTMRVTWAPPSIDLTNFLVRY

SPVKNEEDVAELSIISPSDNAVVLTNLLPGTEYVVSVSS

SUPPLEMENTAL PRELIMINARY AMENDMENT AND TRANSMITTAL OF SEQUENCE LISTING

VYEQHESTPLRGRQKTGLDSPTGIDFSIDTANSFTVHW

IAPRATITGYRIRHHPEHFSGRPREDRVPHSRNSITLT

NLTPGTEYVVSIVALNGREESPLLIGRSRSRSHHHHH"

[SEQ ID NO:32] [SEQ ID NO:27]

Note1: Residue 1 to 207 is pQE sequence from and including Qiagen promoter primer (CCCGAAAAGTGCCACCTG) [SEQ ID N:29]. Residue 1342 to 1399 is pQE12 sequence from the end of the hexa-histidine tag to the Qiagen reverse primer sequence (GTTCTGAGGTCAATTACTGG) [SEQ ID NO:30]. Fibronectin-derived sequence (*i.e.* without MRGS and hexa-histidine tag is in lower case).

Please replace the paragraph on page 48, lines 5-38 with the following paragraph.

BASE COUNT 390 a 368 c 290 g 351 t  
ORIGIN

1 CCCCGAAAAG TGCCACCTGA CGTCTAAGAA ACCATTATTA TCATGACATT  
AACCTATAAA

61 AATAGGCGTA TCACGAGGCC CTTTCGTCTT CACCTCGAGA AATCATAAAA  
AATTTATTTG

121 CTTTGTGAGC GGATAACAAT TATAATAGAT TCAATTGTGA GCGGATAACA  
ATTCACACA

181 GAATTCATTA AAGAGGAGAA ATTAACATATG AGAGGATCtg tggtgacacc  
attgtctcca

241 ccaacaaact tgcatctgga ggcaaaccct gacactggag tgctcacagt ctcctggag

301 aggagcacca ccccagacat tactggttat agaattacca caaccctac aaacggccag

SUPPLEMENTAL PRELIMINARY AMENDMENT AND TRANSMITTAL OF SEQUENCE LISTING

361 caggaaatt ctggaaaga agtggccat gctgatcaga gtcctgcac tttgataac  
421 ctgagtcccg gcctggagta caatgtcagt gtttacactg tcaaggatga caaggaaagt  
481 gtccttatct ctgataccat catcccagag gtgcaccaac tcactgacct aagcttgtt  
541 gatataaccg attcaagcat cggcctgagg tggacccgc taaacttttc caccattatt  
601 gggtaccgca tcacagtagt tgcggcagga gaaggatcc ctattttga agattttgt  
661 gactcctcag taggatacta cacagtcaca gggctggagc cggcattga ctatgatatc  
721 agcgttatca ctctcattaa tggccggcag agtgcctcta ctacactgac acaacaaacg  
781 gctgttcctc ctcccactga cctgcgattc accaacattg gtccagacac catcggtgc  
841 acctgggctc caccatcattt accaacttcc tggtgcgtta ctacactgt  
901 aaaaatgagg aagatgtgc agagtttca atttctccctt cagacaatgc agtggcttta  
961 acaaatctcc tgccgttac agaatatgta gtgagtgctt ccagtgtcta cgaacaacat  
1021 gagagcacac ctcttagagg aagacagaaa acaggtctt attccccac tggcattgac  
1081 ttctgtata ttactgccaa ctctttact gtgcactgga ttgctctcg agccaccatc  
1141 actggctaca ggatccgcca tcatcccgag cacttcagtg ggagacctcg agaagatcgg  
1201 gtgccttact ctccggattc catcaccctc accaacctca ctccaggcac agagttatgt  
1261 gtcagcatcg ttgctctaa tggcagagag gaaagtccct tattgattgg caGATCCAGA  
1321 TCTCATCACC ATCACCATCA CTAAGCTTAA TTAGCTGAGC TTGGACTCCT  
  
GTTGATAGAT  
  
1381 CCAGTAATGA CCTCAGAAC

[SEQ ID NO:33] [SEQ ID NO: 28]

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